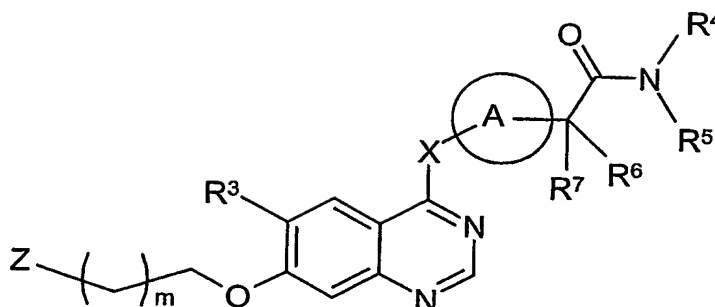


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CLAIMS

1. A compound of formula (I):



formula (I)

wherein A is 5-membered heteroaryl containing a nitrogen atom and optionally containing one or two further nitrogen atoms;

X is O, S, S(O), S(O)₂ or NR¹⁴;

m is 0, 1, 2 or 3;

10 Z is a group selected from -NR¹R², phosphonooxy, C₃₋₆cycloalkyl which C₃₋₆cycloalkyl is substituted by phosphonooxy or C₁₋₄alkyl substituted by phosphonooxy, and a 4- to 7-membered ring linked via a carbon atom containing a nitrogen atom and optionally containing a further nitrogen atom, which ring may be saturated, partially saturated or unsaturated wherein the ring is substituted on carbon or nitrogen by phosphonooxy or C₁₋₄alkyl

15 substituted by phosphonooxy, and wherein the ring is optionally further substituted on carbon or nitrogen by 1, 2 or 3 halo or C₁₋₄alkyl groups;

R¹ is a group selected from -COR⁸, -CONR⁸R⁹ and C₁₋₆alkyl which C₁₋₆alkyl is substituted by phosphonooxy and optionally further substituted by 1 or 2 halo or methoxy groups;

20 R² is a group selected from hydrogen, -COR¹⁰, -CONR¹⁰R¹¹ and C₁₋₆alkyl which C₁₋₆alkyl is optionally substituted by 1, 2 or 3 halo or C₁₋₄alkoxy groups or -S(O)_pR¹¹ (where p is 0, 1 or 2) or phosphonooxy, or R² is a group selected from C₂₋₆alkenyl, C₂₋₆alkynyl, C₃₋₆cycloalkyl and C₃₋₆cycloalkylC₁₋₄alkyl;

or R¹ and R² together with the nitrogen to which they are attached form a 4- to 7- membered ring optionally containing a further nitrogen atom which ring may be saturated, unsaturated or 25 partially saturated wherein the ring is substituted on carbon or nitrogen by a group selected from phosphonooxy and C₁₋₄alkyl which C₁₋₄alkyl is substituted by phosphonooxy or -NR⁸R⁹, and where the ring is optionally further substituted on carbon or nitrogen by 1, 2 or 3 halo or C₁₋₄alkyl groups;

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R^3 is a group selected from hydrogen, halo, cyano, nitro, C_{1-6} alkoxy, C_{1-6} alkyl, $-OR^{12}$, $-CHR^{12}R^{13}$, $-OC(O)R^{12}$, $-C(O)R^{12}$, $-NR^{12}C(O)R^{13}$, $-C(O)NR^{12}R^{13}$, $-NR^{12}SO_2R^{13}$ and $-NR^{12}R^{13}$;

R^4 is hydrogen or a group selected from C_{1-4} alkyl, heteroaryl, heteroaryl C_{1-4} alkyl, aryl and aryl C_{1-4} alkyl which group is optionally substituted by 1, 2 or 3 substituents selected from halo, methyl, ethyl, cyclopropyl and ethynyl;

R^5 is selected from hydrogen, C_{1-4} alkyl, C_{2-4} alkenyl, C_{2-4} alkynyl, C_{3-6} cycloalkyl and C_{3-6} cycloalkyl C_{1-4} alkyl;

R^6 and R^7 are independently selected from hydrogen, halo, C_{1-4} alkyl, C_{3-6} cycloalkyl, hydroxy and C_{1-4} alkoxy;

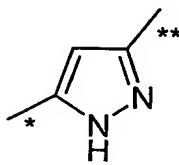
R^8 is C_{1-4} alkyl substituted by phosphonooxy and optionally further substituted by 1 or 2 halo or methoxy groups;

R^9 is selected from hydrogen and C_{1-4} alkyl;

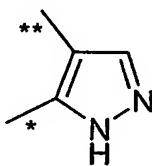
R^{10} is selected from hydrogen and C_{1-4} alkyl (optionally substituted by halo, C_{1-4} alkoxy, $S(O)_q$ (where q is 0, 1 or 2) or phosphonooxy);

R^{11} , R^{12} , R^{13} and R^{14} are independently selected from hydrogen, C_{1-4} alkyl and heterocyclyl; or a pharmaceutically acceptable salt thereof.

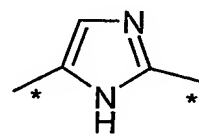
2. A compound according to claim 1 wherein A is a group of formula (a), (b), (c), (d) or (e):



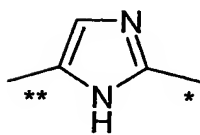
(a)



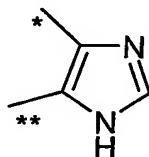
(b)



(c)



(d)



(e)

where * is the point of attachment to the X group of formula (I) and ** is the point of attachment to the (CR^6R^7) group of formula (I); or a pharmaceutically acceptable salt thereof.

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3. A compound according to claim 2 wherein A is a group of formula (a) as defined in claim 2; or a pharmaceutically acceptable salt thereof.
- 5 4. A compounds according to any one of claims 1, 2 or 3 wherein X is NH; or a pharmaceutically acceptable salt thereof.
5. A compound according to any one of the preceding claims wherein Z is $-NR^1R^2$ or a 5- to 6-membered saturated ring linked via a carbon atom containing a nitrogen atom and
10 optionally containing a further nitrogen atom, wherein the ring is substituted on carbon or nitrogen by phosphonooxy or C_{1-4} alkyl substituted by phosphonooxy; or a pharmaceutically acceptable salt thereof.
6. A compound according to any one of the preceding claims wherein R^1 is C_{1-5} alkyl
15 substituted by phosphonooxy and R^2 is a group selected from hydrogen and C_{1-6} alkyl which C_{1-6} alkyl is optionally substituted by 1, 2 or 3 halo or C_{1-4} alkoxy groups, or R^2 is a group selected from C_{2-6} alkenyl, C_{2-6} alkynyl, C_{3-6} cycloalkyl and C_{3-6} cycloalkyl C_{1-4} alkyl; or a pharmaceutically acceptable salt thereof.
- 20 7. A compound according to any one of the preceding claims wherein R^1 is 2-phosphonooxyethyl; or a pharmaceutically acceptable salt thereof.
8. A compound according to any one of claims 1 to 5 where Z is $-NR^1R^2$ and R^1 and R^2 together with the nitrogen to which they are attached form a piperidine, pyrrolidine or
25 piperazine ring which is substituted by a group selected from phosphonooxy, phosphonooxymethyl, 2-phosphonooxyethyl, *N*-ethyl-*N*-(2-phosphonooxyethyl)aminomethyl and *N*-(2-phosphonooxyethyl)aminomethyl and where the ring is optionally further substituted by 1 or 2 methyl; or a pharmaceutically acceptable salt thereof.
- 30 9. A compound according to claim 8 wherein R^1 and R^2 together with the nitrogen to which they are attached form 2-(phosphonooxymethyl)pyrrolidinyl; or a pharmaceutically acceptable salt thereof.

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10. A compound according to any one of the preceding claims wherein R⁴ is 3-fluorophenyl, 3,5-difluorophenyl or 2,3-difluorophenyl; or a pharmaceutically acceptable salt thereof.
- 5 11. A compound according to any one of the preceding claims wherein R³ is C₁₋₄alkoxy, halo or hydrogen; or a pharmaceutically acceptable salt thereof.
12. A compound selected from:
- 1-[3-({4-[(5-{2-[(3-fluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-6-methoxyquinazolin-7-yl}oxy)propyl]piperidin-4-yl}methyl dihydrogen phosphate;
- 10 2-[[3-({4-[(5-{2-[(3,5-difluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-6-methoxyquinazolin-7-yl}oxy)propyl](ethyl)amino]ethyl dihydrogen phosphate;
- {(2*S*)-1-[3-({4-[(5-{2-[(3,5-difluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-6-methoxyquinazolin-7-yl}oxy)propyl]pyrrolidin-2-yl}methyl dihydrogen phosphate;
- 15 {(2*R*)-1-[3-({4-[(5-{2-[(3,5-difluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-6-methoxyquinazolin-7-yl}oxy)propyl]pyrrolidin-2-yl}methyl dihydrogen phosphate;
- {(2*S*)-1-[3-({4-[(5-{2-[(3-fluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-6-methoxyquinazolin-7-yl}oxy)propyl]pyrrolidin-2-yl}methyl dihydrogen phosphate;
- 20 2-[[3-({4-[(5-{2-[(2,3-difluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-6-methoxyquinazolin-7-yl}oxy)propyl](propyl)amino]ethyl dihydrogen phosphate;
- 2-[[3-({4-[(5-{2-[(2,3-difluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-6-methoxyquinazolin-7-yl}oxy)propyl](isobutyl)amino]ethyl dihydrogen phosphate;
- 2-[[3-({4-[(5-{2-[(3,5-difluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-6-methoxyquinazolin-7-yl}oxy)propyl](isobutyl)amino]ethyl dihydrogen phosphate;
- 25 2-[[3-({4-[(5-{2-[(3,5-difluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-6-methoxyquinazolin-7-yl}oxy)propyl](propyl)amino]ethyl dihydrogen phosphate;
- 2-[[3-({4-[(5-{2-[(3-fluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-6-methoxyquinazolin-7-yl}oxy)propyl](isobutyl)amino]ethyl dihydrogen phosphate;
- 2-[(2,2-dimethylpropyl)[3-({4-[(5-{2-[(3-fluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-6-methoxyquinazolin-7-yl}oxy)propyl]amino]ethyl dihydrogen phosphate;
- 30 1-[3-({4-[(5-{2-[(3-fluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-6-methoxyquinazolin-7-yl}oxy)propyl]piperidin-3-yl dihydrogen phosphate;

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- {(2*R*)-1-[3-(4-[(5-{2-[(2,3-difluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-6-methoxyquinazolin-7-yl}oxy)propyl]pyrrolidin-2-yl}methyl dihydrogen phosphate;
- 2-[[3-(4-[(5-{2-[(3,5-difluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-6-methoxyquinazolin-7-yl}oxy)propyl](prop-2-yn-1-yl)amino]ethyl dihydrogen phosphate;
- 5 2-[[3-(4-[(5-{2-[(2,3-difluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-6-methoxyquinazolin-7-yl}oxy)propyl](isopropyl)amino]ethyl dihydrogen phosphate;
- 2-[[3-(4-[(5-{2-[(2,3-difluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-6-methoxyquinazolin-7-yl}oxy)propyl](prop-2-yn-1-yl)amino]ethyl dihydrogen phosphate;
- 2-[[3-(4-[(5-{2-[(2,3-difluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-6-methoxyquinazolin-7-yl}oxy)propyl](2-methoxyethyl)amino]ethyl dihydrogen phosphate;
- 10 2-[[3-(4-[(5-{2-[(3-fluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-6-methoxyquinazolin-7-yl}oxy)propyl]amino}ethyl dihydrogen phosphate;
- 2-[(cyclobutylmethyl)[3-(4-[(5-{2-[(2,3-difluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-6-methoxyquinazolin-7-yl}oxy)propyl]amino}ethyl dihydrogen phosphate;
- 15 2-[[3-(4-[(5-{2-[(3-fluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-6-methoxyquinazolin-7-yl}oxy)propyl](3,3,3-trifluoropropyl)amino]ethyl dihydrogen phosphate;
- 2-{allyl[3-(4-[(5-{2-[(2,3-difluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-6-methoxyquinazolin-7-yl}oxy)propyl]amino}ethyl dihydrogen phosphate;
- 20 2-{cyclobutyl[3-(4-[(5-{2-[(2,3-difluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-6-methoxyquinazolin-7-yl}oxy)propyl]amino}ethyl dihydrogen phosphate;
- 2-{cyclopentyl[3-(4-[(5-{2-[(2,3-difluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-6-methoxyquinazolin-7-yl}oxy)propyl]amino}ethyl dihydrogen phosphate;
- 2-{cyclopropyl[3-(4-[(5-{2-[(3-fluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-6-methoxyquinazolin-7-yl}oxy)propyl]amino}ethyl dihydrogen phosphate;
- 25 2-{(cyclopropylmethyl)[3-(4-[(5-{2-[(2,3-difluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-6-methoxyquinazolin-7-yl}oxy)propyl]amino}ethyl dihydrogen phosphate;
- 2-{cyclobutyl[3-(4-[(5-{2-[(3-fluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-6-methoxyquinazolin-7-yl}oxy)propyl]amino}ethyl dihydrogen phosphate;
- 30 2-{4-[(4-[(5-{2-[(2,3-difluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]quinazolin-7-yl}oxy)methyl]piperidin-1-yl}ethyl dihydrogen phosphate;
- 2-[[3-(4-[(5-{2-[(2,3-difluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-quinazolin-7-yl}oxy)propyl](ethyl)amino]ethyl dihydrogen phosphate;

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- 2-[[3-(4-[(5-{2-[(2,3-difluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-quinazolin-7-yl}oxy)propyl](isopropyl)amino]ethyl dihydrogen phosphate;
- 3-[[3-(4-[(5-{2-[(2,3-difluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]quinazolin-7-yl}oxy)propyl]amino]-3-methylbutyl dihydrogen phosphate;
- 5 2-[(2*S*)-1-[3-(4-[(5-{2-[(2,3-difluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-quinazolin-7-yl}oxy)propyl]pyrrolidin-2-yl}ethyl dihydrogen phosphate;
- {(2*R*)-1-[3-(4-[(5-{2-[(2,3-difluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-quinazolin-7-yl}oxy)propyl]pyrrolidin-2-yl}methyl dihydrogen phosphate;
- 2-[[3-(4-[(5-{2-[(2,3-difluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-quinazolin-7-yl}oxy)propyl](propyl)amino]ethyl dihydrogen phosphate;
- 10 2-[[3-(4-[(5-{2-[(2,3-difluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-quinazolin-7-yl}oxy)propyl](butyl)amino]ethyl dihydrogen phosphate;
- 2-{cyclopentyl[3-(4-[(5-{2-[(2,3-difluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-quinazolin-7-yl}oxy)propyl]amino}ethyl dihydrogen phosphate;
- 15 {(2*S*)-1-[3-(4-[(5-{2-[(2,3-difluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-quinazolin-7-yl}oxy)propyl]pyrrolidin-2-yl}methyl dihydrogen phosphate;
- {(2*S*)-1-[3-(4-[(5-{2-[(3-fluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-quinazolin-7-yl}oxy)propyl]pyrrolidin-2-yl}methyl dihydrogen phosphate;
- 2-{cyclopentyl[3-(4-[(5-{2-[(3-fluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-quinazolin-7-yl}oxy)propyl]amino}ethyl dihydrogen phosphate;
- 20 2-[[3-(4-[(5-{2-[(3-fluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-quinazolin-7-yl}oxy)propyl](ethyl)amino]ethyl dihydrogen phosphate;
- 2-[[3-(4-[(5-{2-[(3-fluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]quinazolin-7-yl}oxy)propyl]amino]-2-methylpropyl dihydrogen phosphate;
- 25 2-[[3-(4-[(5-{2-[(3-fluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-quinazolin-7-yl}oxy)propyl](propyl)amino]ethyl dihydrogen phosphate;
- {(2*R*)-1-[3-(4-[(5-{2-[(3-fluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-quinazolin-7-yl}oxy)propyl]pyrrolidin-2-yl}methyl dihydrogen phosphate;
- 3-[[3-(4-[(5-{2-[(3-fluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-quinazolin-7-yl}oxy)propyl](ethyl)amino]propyl dihydrogen phosphate
- 30 2-[[3-(4-[(5-{2-[(3-fluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]quinazolin-7-yl}oxy)propyl](2-methoxyethyl)amino]ethyl dihydrogen phosphate

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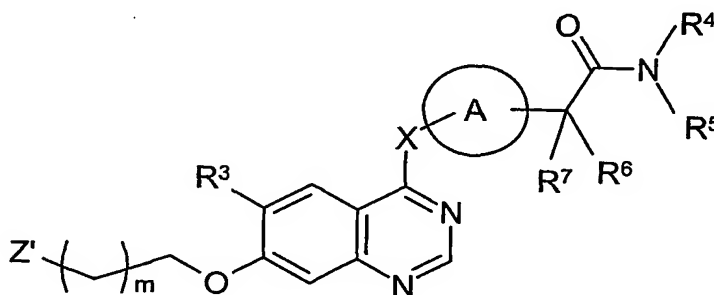
- 2-[[4-(4-[(5-{2-[(2,3-difluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-quinazolin-7-yl}oxy)butyl](propyl)amino]ethyl dihydrogen phosphate;
- 2-[[4-(4-[(5-{2-[(2,3-difluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-quinazolin-7-yl}oxy)butyl](ethyl)amino]ethyl dihydrogen phosphate;
- 5 {(2*R*)-1-[4-(4-[(5-{2-[(2,3-difluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-quinazolin-7-yl}oxy)butyl]pyrrolidin-2-yl}methyl dihydrogen phosphate;
- 2-[[4-(4-[(5-{2-[(2,3-difluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]quinazolin-7-yl}oxy)butyl](methyl)amino]ethyl dihydrogen phosphate;
- {(2*S*)-1-[4-(4-[(5-{2-[(2,3-difluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]-
- 10 quinazolin-7-yl}oxy)butyl]pyrrolidin-2-yl}methyl dihydrogen phosphate; and
- 2-{ethyl[3-(6-fluoro-4-[(5-{2-[(3-fluorophenyl)amino]-2-oxoethyl}-1*H*-pyrazol-3-yl)amino]quinazolin-7-yl}oxy)propyl]amino}ethyl dihydrogen phosphate;
- or a pharmaceutically acceptable salt thereof.
- 15 13. A pharmaceutical composition comprising a compound according to any one of the preceding claims in association with a pharmaceutically acceptable diluent or carrier.
14. Use of a compound according to any one of claims 1 to 12 in therapy.
- 20 15. Use of a compound according to any one of claims 1 to 12 in the preparation of a medicament for the treatment of a disease where the inhibition of one or more Aurora kinase is beneficial.
16. Use according to claim 15 wherein Aurora kinase is Aurora-A kinase or Aurora-B
- 25 kinase.
17. Use of a compound according to any one of claims 1 to 12 in the preparation of a medicament for the treatment of colorectal, breast, lung, prostate, pancreatic or bladder and renal cancer or leukemias or lymphomas
- 30
18. A method of treating a human suffering from a disease in which the inhibition of one or more Aurora kinases is beneficial, comprising the steps of administering to a person in need thereof a therapeutically effective amount of a compound as defined in claim 1 or a

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pharmaceutically acceptable salt thereof.

19. A method of treating a human suffering from colorectal, breast, lung, prostate, pancreatic or bladder and renal cancer or leukemias or lymphomas, comprising the steps of
 5 administering to a person in need thereof a therapeutically effective amount of a compound as defined in claim 1 or a pharmaceutically acceptable salt thereof.

20. A process for the preparation of a compound of formula (I) as defined in claim 1 or a pharmaceutically acceptable salt thereof, which process comprises converting a compound of
 10 formula (II) into a compound of formula (I) by phosphorylation of an appropriate hydroxy group:



formula (II)

where A, X, m, R³, R⁴, R⁵, R⁶, R⁷ and R⁹ are as defined for formula (I); and Z' is a group
 15 selected from -NR^{1'}R^{2'}, hydroxy, C₃₋₆cycloalkyl which C₃₋₆cycloalkyl is substituted by hydroxy or C₁₋₄alkyl substituted by hydroxy, and a 4- to 7-membered ring linked via a carbon atom containing a nitrogen atom and optionally containing a further nitrogen atom, which ring may be saturated, unsaturated or partially saturated wherein the ring is substituted on carbon or nitrogen by hydroxy or C₁₋₄alkyl substituted by hydroxy and wherein the ring is optionally
 20 further substituted on carbon or nitrogen by 1, 2 or 3 halo or C₁₋₄alkyl groups; R^{1'} is a group selected from -COR^{8'}, -CONR^{8'}R⁹ and C₁₋₆alkyl which C₁₋₆alkyl is substituted by hydroxy and optionally further substituted by 1 or 2 halo or methoxy groups; R^{2'} is a group selected from hydrogen, -COR¹⁰, -CONR¹⁰R¹¹ and C₁₋₆alkyl which C₁₋₆alkyl is optionally substituted by 1, 2 or 3 halo or C₁₋₄alkoxy groups or -S(O)_pR¹¹ (where p is 0, 1 or 2) or hydroxy, or R² is
 25 a group selected from C₂₋₆alkenyl, C₂₋₆alkynyl, C₃₋₆cycloalkyl and C₃₋₆cycloalkylC₁₋₄alkyl; or R^{1'} and R^{2'} together with the nitrogen to which they are attached form a 4- to 7- membered ring optionally containing a further nitrogen atom which may be saturated, unsaturated or

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partially saturated wherein the ring is substituted on carbon or nitrogen by a group selected from hydroxy and C₁₋₄alkyl which C₁₋₄alkyl is substituted by hydroxy or -NR^{8'}R⁹ and where the ring is optionally further substituted on carbon or nitrogen by 1, 2 or 3 halo or C₁₋₄alkyl groups; and where R^{8'} is C₁₋₄alkyl substituted by hydroxy and optionally further substituted

5 by 1 or 2 halo or methoxy groups:

and thereafter if necessary:

i) converting a compound of formula (I) into another compound of formula (I); and/or

ii) removing any protecting groups; and/or

iii) forming a pharmaceutically acceptable salt thereof.